(51) International Patent Classification7: C07D 207/46, 295/24

- (21) International Application Number: PCT/BR00/00068
- 21 June 2000 (21.06.2000) (22) International Filing Date:
- (25) Filing Language:

English

(26) Publication Language;

English

(30) Priority Data: PI 9903137-0 1

22 June 1999 (22.06.1999)

- (71) Applicant (for all designated States except US): CON-SELHO NACIONAL DE DESENVOLVIMENTO CIENTÍFICO E TECNOLÓGICO - CNPQ [BR/BR]; Avenida W3 Norte, Quadra 507/B, Lote 2, SEPN, CEP-70741-901 Brasilia, DF (BR).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): NAKAIE, Clovis, Ryuichi [BR/BR]; CEP-São Paulo, SP (BR).

TOMINAGA, Mineko [BR/BR]; CEP-São Paulo, SP (BR). DE MATTOS PAIVA, Antonio, Cechelli [BR/BR]; CEP-São Paulo, SP (BR). DOS REIS BAR-BOSA, Simone [BR/BR]; CEP-São Paulo, SP (BR). MARCHETTO, Reinaldo [BR/BR]; CEP-São Paulo, SP (BR). SCHREIER, Shirley [BR/BR]; CEP-São Paulo, SP

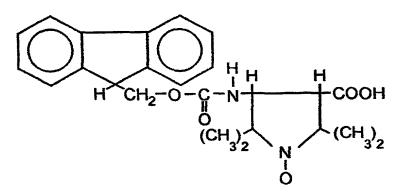
- EURY PEREIRA, Luna-Filho; Avenida (74) Agent: W3 Norte, Quadra 507/B, Lote 2, SEPN, Suite 211, CEP-70741-901 Brasilia, DF (BR).
- (81) Designated States (national): JP, US.
- (84) Designated States (regional): European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).

Published:

- With international search report.
- Before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments.

[Continued on next page]

(54) Title: SYNTHESIS OF A NOVEL PARAMAGNETIC AMINO ACID DERIVATIVE (EPM-5) FOR LABELLING CHEMI-CAL AND BIOLOGICAL MACROMOLECULES



Structure of Fmoc-Poac.

(57) Abstract: The present invention refers to the synthesis and application of 2,2,5,5-tetramethylpyrrolidine-N-oxyl-(9-fluorenylmethyloxycarbonyl)-3-amine-4-carboxylic acid, a novel paramagnetic (spin label) amino acid derivative denominated as Fmoc-POAC whose structure is seen in Figure 1. Fmoc-POAC can be coupled to peptide sequences and other systems. It can be inserted anywhere in a peptide segment, even at an internal position, after removal of its temporary amine protecting group, Fmoc. Owing to its pyrrolidine structure, this molecule may induce differentiated conformations if compared with the normal alpha-amino acids, thus becoming a valuable probe for strctural-biological activity of certain peptides. The POAC-angiotensin II analogue was synthesized as a model according to the use of the chemical derivative.